

10 30 50 70
 gaattccggcgagtgaggcgctgacagggaactcggggggactcttgacagagcccttgaccacgccgccatcgagcctccag
 90 110 130 150 170
 ccagtcctctctgcccgttctctcgccatggaggccgcccgcgtccgagggttcgagcagcgagccggcgaggct
 190 210 230 250
 gaccccolgtggcgagggcccggtcctggaggcggagctgccgtgcgctccccgcgggtcccccccgagcgcgggctcgggt
 270 290 310 330
 cagcatggattcctgggttcatttctgtccctglttgccagtggtctaatcatglttagtgccacccatgctactacagtttcacc
 350 370 390 410
 M D S W F I L V L F G S G L I H V S A N N A T T V S P
 ttctttgggaacgacagatttaataaacatcaacacaggaattggctaaaggaggaataaacctcaaatlcaacctcttc
 430 450 470 490
 S L G T T R L I K T S T T E L A K E E N K T S N S T S S
 agtaatttctcttctglggaccccaacttcagcccaacttgactctggagcccaactatgtagctactgttaatttctcaco
 510 530 550 570 590
 V I S L S V A P T F S P N L T L E P T Y V T T V N S S H
 ctctgacaatgggocccaggaggcgccagcagcgggaactcggggcactaccatttccccgagcgggaactggttattgagaa
 610 630 650 670
 S D N G T R R A A S T E S G G T T I S P N G S W L I E N
 ccagltcacggtgccatcaacagaacccctgggggggaactccagcactgcagcaaccactccagaaacttccccccggcaga
 690 710 730 750
 Q F T D A I T E P W E G N S S T A A T T P E T F P P A D
 lgagoccccaattattgcggtgatggcgccctgctctctgctgtagtaactggttattatcatagltctgtacatgttaag
 770 790 810 830
 E T P I I A V M V A L S S L L V I V F I I I V L Y M L R
 gtttaagaaatcaacgcaagctgggggtcatttcccaactcttccgcctgltcaaatggccgcagcggggtgtgggcccccaag
 F K K Y K Q A G S H S N S F R L S N G R T E D V E P Q S

FIG.1A

FIG. 1B

1610 1630 1650 1670
 ccctgctggacatgatgcattcggagcgcaaaagtgatgtataggggttgtagccggatccgggcccgctgccagatggtg
 M L D M M H S E R K V D V Y G F V S R I R A Q R C Q M V
 1690 1710 1730 1750 1770
 cagacagacatgcagtcgctcctcatataccaggcccttcggagcattatctgtatggggccacagaaactgggaagtgacttctc
 Q T D M Q Y V F I Y] Q A L L E H Y L Y G D T E L E V T S L
 1790 1810 1830 1850
 / tggaaaccacctacaaaaatttataoacagatccacggactagcaaacccgggttagaggagggtltaagaaatlaacttc
 E T H L Q K I Y N K I P G T S N N G L E E F K K L T S
 1870 1890 1910 1930
 ootcaaaatccagaatgacaagatgcgcacggggaaccltccagccaacatgaaggaaggaacccgggtllacagatcaltccat
 I K I Q N D K M R T G N L P A N M K [K N R V L Q I I P Y
 1950 1970 1990 2010
 gaatttaacagagtgatcattccagtcacacggggaaggaaggaacacagactatgtgaacgcaltcaltgatggatccggc
 E F N R V I I P V K R G E E N T D Y V N A S F I D C Y R Q
 2030 2050 2070 2090 2110
 agaaagactcctacattgccagccaggccctcttctccacacgatggagacttctggcgaatgatctgggagtggaagtccctg
 K D S Y I A S Q G P L L H T I E D F W R M I W E W K S C
 2130 2150 2170 2190
 ttctatcgtaatgctgacagaaactggaaagagagggccaggaagtgtagccagtaactggccatctgatggcctgggtgctctac
 S I V M L T E L E E R G Q E K C A Q Y W P S D G L V S Y
 2210 2230 2250 2270
 ggagocacagttgagctgaaggaaggaaggaatgtgaagactacactgtccggagacctctggtcaccacaccaggagga II
 G D I T V E L K K E E C E S Y T V R D L L V T N T R E N

FIG.1C

0.1 kb

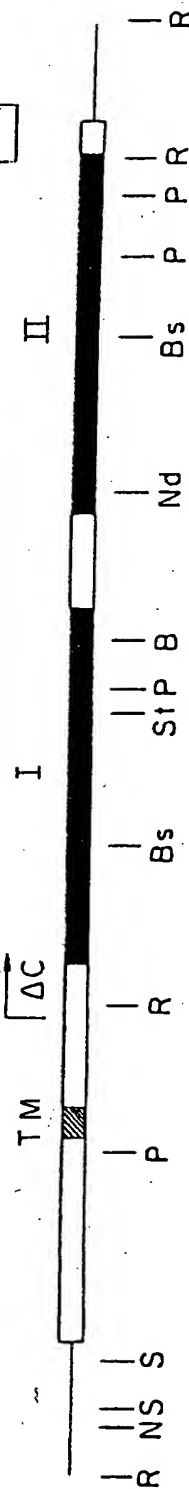


FIG. 1E

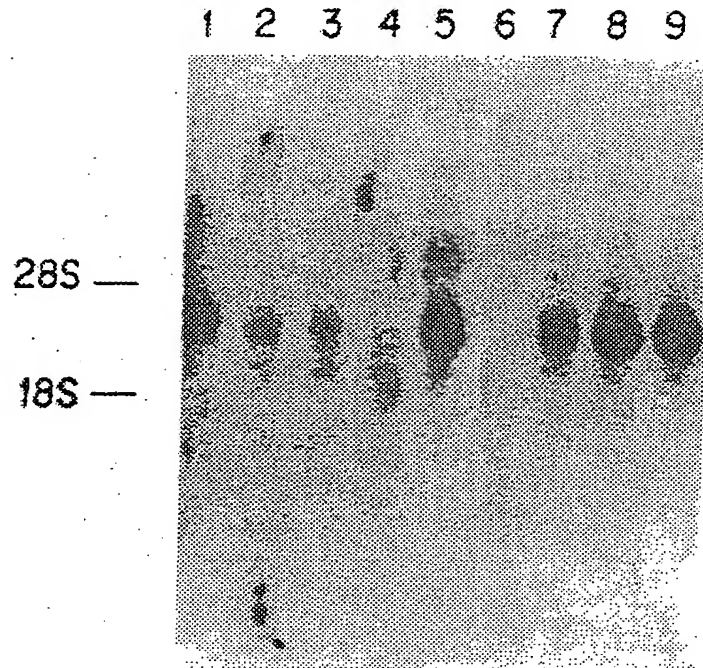


FIG. 2

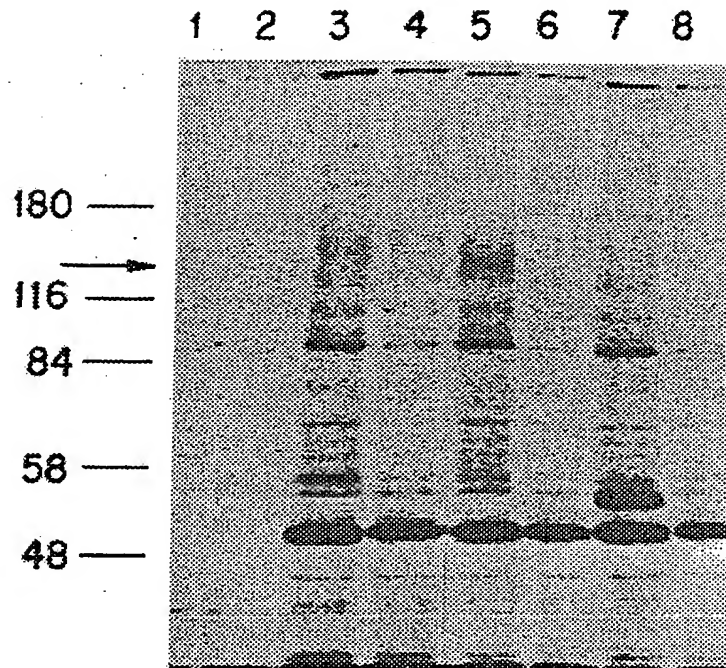
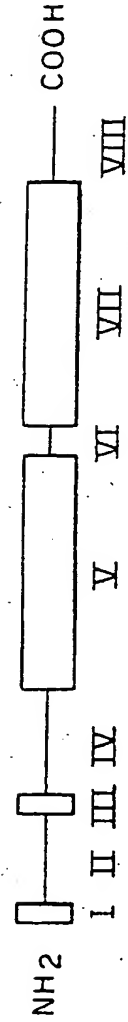


FIG. 3



①
 1 MDSWF ILVLLGSLICVS ANNATTVAPSVGI IRLINSSTAEPVKEEAKTSNPTSSLISLSVAPTFSF 68
 2 F H S L T K T I L A N S V I 68
 1 NITLGPTYLT TVNSSDSNGITRTASTINSIGITISPNGIWL PDNQFTDARTEPWEGNSSTAATTPEITFP 136
 2 L E V H R A E G T S I E I 136
 1 PSGNSDSKDRRDE TPI IAVMVALSSLLVIVF I I I V L Y M I R F K K Y K Q A G S H S N S F R L S N G R T E D V E P Q S 204
 2 A 195
 1 VPLLARSPSTNRKYPPLPVDKLEEE INRRMADDNKLFREEFNALPACPIQATCEAAASKEENKEKNRYV 272
 2 263
 1 NILPYDHSRVHLTPVEGVDPDSYINASFINGYQEKKNKFAAQPKKEETVNDFWRM IWEQNTAT IVMVT 340
 2 (V) 331
 1 NLKERKECKAQYMPDQGCWTYGNIRVSVEDVTVLVDYTVRKFCIQQVGDMTNRKPQRLITQFHFTSW 408
 2 V S V 399
 1 PDFGVPFTPIGMLKFLKKVKACNPQYAGAIWHCSAGVGRGTGTFWIDAMLDMMHTERKVDVYGVFSR 476
 2 S 467
 1 IRAQRQCMVQTDMMQYVFIYQALLEHLYGYDTELEVTSLETHLQIYNKIPGTSNNGLEEEFKLTSIK 544
 2 535
 1 IQNDKMRITGNLPANMKKNRVLQIIPYEFNRVIIPVKRGEENTIDYVNASFIDGYRQKDSYIASQGPLLH 612
 2 (VII) 603
 1 TIEDFWRM IWEWKSCSIVMLTELEERGQEKCAQYWPDSGLVSYGDI TVELKKEEESYTVRDLLVTN 680
 2 671
 1 TRENKSRQIRQHFHGWPEVGIPSDGKGMISIIAAVQKQQQSGNHPITVHCSAGAGRTGTFCALSTV 748
 2 N 739
 1 LERVKAEGILDVFQTVKSLRLQRPHMVQTLQEQEFCYKVVOEYIDAFSDYANFK 802
 2 793

FIG. 4D

	10	20	30	40	
LCA	NqnKNRYVdILPYDynRVeL	sEinGdagSnYINASyldGfkeEprKyIAA			
RPTase α	NKeKNRYVNILPYDHSRVhLtpvE	GvpdSDYINASfInGYqEknKfIAA			
RPTase β	NKHKNRYINIVAYDHSRVKLaaLaeKDgKltDYINANYVDGYNrpKAYIAA				
RPTase γ	NKHKNRYINIIAYDHSRVKLrpLpgKDsKhsDYINANYVDGYNkaKAYIAA				
CON	NkhKNRY-nII-YDhsRVkL—l—k—k—sdYINA—y—dGynepk—yIAa				
	50	60	70	80	90
LCA	QGPrdETVdDFWRMIWEQkatvIVMVTrceEgnrnKCAeYWPsmEegTra				
RPTase α	QGPkeETVnDFWRMIWEQntatIVMVTNLkErkeckCAQYWPdqGewTYG				
RPTase β	QGPKSTaEDFWRMIWEhNvevIVMITNLVEKGRRKCDQYWPdGSEEEYG				
RPTase γ	QGPKSTfEDFWRMIWEqntgiIVMITNLVEKGRRKCDQYWPtenSEEEYG				
CON	QGPik-TveDFWRMIWEqnt-vIVM-TnlvEkgrRK-qYWP—gse-yg				
	100	110	120	130	
LCA	fgdVvVkinqhkrpcDYiiqKI	nIvn	kkekatgRevThiq		
RPTase α	NirVsVedVtVLv	DYTVRKFc	IqqvGd	mtnRkpqRIiTQfH	
RPTase β	NfIVTqKSVqVLA	yYTVRnFtIRNTKIKK	Gs	qKGRpsgRVVTQYH	
RPTase γ	NiIVTIKStkihAc	YTVRrFsIRNTKvKK	GqkgnpKGRqneRVViQYH		
CON	ni-Vlvk-v-vla—dYlvrkf—rntki—k—g—k—kgr—gRvvtqyh				

FIG.5A

	140	150	160	170	180	190
LCA	FTSWPDhGV	PedPhlILKlrrr	VnAfsnffs	GpIVVHCSAGVGRTGT	YigID	
RPTase α	FTSWPDfGV	PftPi gmlKF l kKVkAcnpqya	GaIVVHCSAGVGRTGT	f vVID		
RPTase β	YTQWPDmGV	PEYsLPVL TFVRK	aayAkrhavGPVVHCSAGVGRTGT	YIVID		
RPTase γ	YTQWPDmGV	PEYdLPVL TFVRr	ssaArmpetGPVIVHCSAGVGRTGT	YIViD		
CON	-T-WPDmGV	PeyplpvL-fvr-v-aa-	Gp-vVHCSAGVGRTGT	YiVID		
	200	210	220	230		
LCA	AMLegl eaEnKVDVY	GyVvkIRrQRCIMVQ	veaQYi l l hQALvE			
RPTase α	AMLdmmhtErKVDVY	GFVsrIRaQRCqMVQTdmQYVF	lyQALIE			
RPTase β	SMLQQIqhEgTVNi	fGFLKHIRsQRNYLVQTEE	QYVF IHDtLvE			
RPTase γ	SMLQQIkdksTVNv	IGFLKHIRtQRNYLVQTEE	QYi f IHDaLIE			
CON	-MLqqi-e-V-vyGf-kh	iR-QR-y-VQteeQY-f	lh-aL-E			

FIG. 5B

	10	20	30	40
LCA	NksKNRnsnvIPYdyNRVp	lkhelemskesehdssdessddds	EEpskY	
RPTase α	NmkKNRvIqIIPYEfNRVilpvkr		GEEenTDY	
RPTase β	NrEKNRLSSIIPvERsRVGIssLs		GE GTDY	
RPTase γ	NkEKNRnSSvvPsERaRVGIapLp		GmkGTDY	
CON	NkeKNRnss-iPyernRVg	l	geegtdY	
	50	60	70	80
LCA	iNASFImSYwkpevmIAaQGPLkeTlgDFWqMI	fqrKvkVIMLTELkhg		
RPTase α	vNASFIdGYrQkdsyIAaQGPLLHTleDFWRMI	WewKscsIMLTELeer		
RPTase β	INASYIMGYYSNEFIITQHPLLHTIKDFWRMI	WOHNAQIVVMIPDgQnm		
RPTase γ	INASYIMGYYSNEFIITQHPLpHTIKDFWRMI	WOHNAQIVVMIPDnQsI		
CON	iNAS-ImGYysnefIITQ-PLIhTikDFWRMI	wdh-naqiVMI-q		
	100	110	120	130
LCA	dQEiCAQYWgeGkqtYGDleVdLKdtdksstYTI	RvfeIrhskrkdsRrtv		
RPTase α	gQEiCAQYWPsdGlvSYGDItVeLKkeeeCESYTV	RdIlvtntreNkSRqI		
RPTase β	A EDEFVYWPn	kDEpi	NCESFkVTLmaeehkCLSNEEkII	
RPTase γ	A EDEFVYWPs	reEsm	NCEaFtVTLiskdrICLSNEEqil	
CON	aE-e-qYWps-g	ygd-v-lk-nces-lvt-e-r-clsne-r-i		
	150	160	170	180
LCA	yQy	qY tnWsvEqIP	aepKellSmIqvVkkQKlpQk	
RPTase α	rQf	HF hgWPevgiP	SdgKgmISilaaV Qk Qq	
RPTase β	IQDFILEATQDDYVLEVRHFQCPKWPNDsPI	SkTFELISVI		K
RPTase γ	IhDFILEATQDDYVLEVRHFQCPKWPNDaPI	SsTFELInVI		K
CON	iqdfileatqddyvlevrhfqcpkwpnpd-Pis-t-ellsvI	qk		

FIG.5C

	190	200	210	220	230
LCA	nsseGNkhhkstP	IIiHCrdGsqqTG	iFCALInI	LEsaetEevvD	iFQvVKa
RPTase α	qqsGNh	PitVHCsaGagr	TGTFCALsTv	LErvkaEgi	IDVFQtVKs
RPTase β	EEAaNR	DGPmIVHDEhGgVt	AGTFCALTTL	mhQLEkENs	VDVyQVAKM
RPTase γ	EEAItR	DGPtIVHDEyGaVs	AGmICALTTL	sQLEnENa	VDVfQVAKM
CON	-eea-nr—	dgP—ivH—e—Gav—	GtFCALttlle	qle—En—vDvfQv—	Km
	240	250			
LCA	LrkaRPgMVstfEQYqFIYdVias				
RPTase α	LaLqRPhMVqTIEQYefcYKVvqe				
RPTase β	INLMRPGVFdIEQYQFIYKVILS				
RPTase γ	INLMRPGVFtIEQYQFIYKarLS				
CON	-nlmRPg—	iEQYqFIYkvils			

FIG.5D

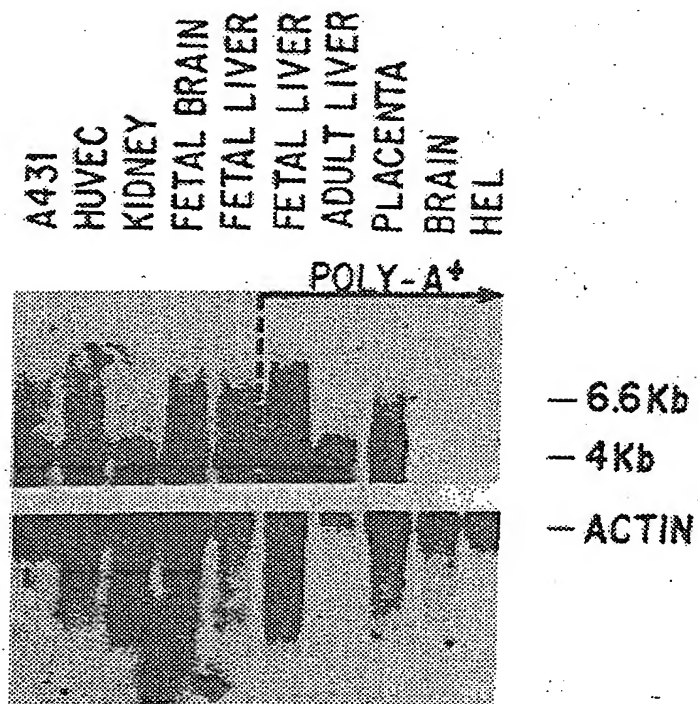
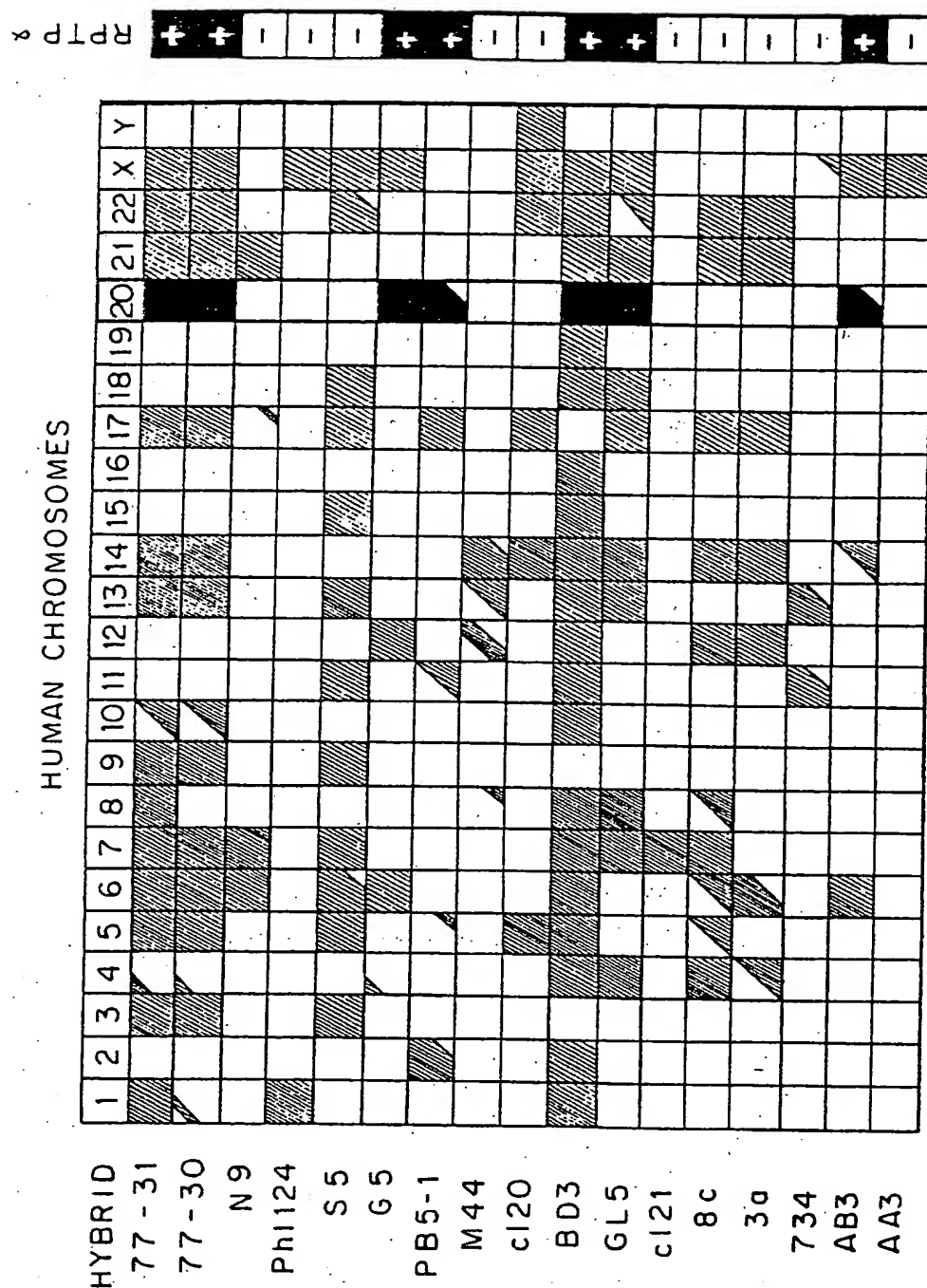


FIG. 6



1 ATGGATTCTGGTTCATTCTTGTTCTGCTCGGCAGTGGTCTGATATGTGTCAGTGCCAAC 60
1[M D S W F I L V L L G S G L I C V S] A N 20
SIGNAL PEPTIDE

61 AATGCTACCACAGTTGCACCTTCTGTAGGAATTACAAGATTAACTCATCAACGGCA 120
21 N A T T V A P S V G I T R L I N S S T A 40

121 GAACCAGTTAAAGAAGAGGCCAAAACTTCAAATCCAACCTTCTTCACTAACTTCTCTTTCT 180
41 E P V K E E A K T S N P T S S L T S L S 60

181 GTGCCACCAACATTCAGCCCAAATATAACTCTGGGACCCACCTATTTAACCACTGTCAAT 240
61 V A P T F S P N I T L G P T Y L T T V N 80

241 TCTTCAGACTCTGACAATGGGACCACAAGAACAGCAAGCACCAATTCTATAGGCATTACA 300
81 S S D S D N G T T R T A S T N S I G I T 100

301 ATTTACCAAATGGAACGTGGCTTCCAGATAACCAGTTCACGGATGCCAGAACAGAACCC 360
101 I S P N G T W L P D N Q F T D A R T E P 120

361 TGGGAGGGGAATTCCAGCACCGCAGCAACCACTCCAGAACTTTCCCTCCTTCAGGTAAT 420
121 W E G N S S T A A T T P E T F P P S G N 140

421 TCTGACTCGAAGGACAGAAGAGATGAGACACCAATTATTGCGGTGATGGTGGCCCTGTCC 480
141 S D S K D R R D E T [P I I A V M V A L S 160
TRANSMEMBRANE SEGMENT

481 TCTCTGCTAGTGATCGTGTTTATTATCATAGTTTGTACATGTTAAGGTTTAAGAAATAC 540
161 S L L V I V F I I I V L Y M L] R F K K Y 180

541 AAGCAAGCTGGGAGCCATTCCAATTCTTTCCGCTTATCCAACGGCCGCACTGAGGATGTG 600
181 K Q A G S H S N S F R L S N G R T E D V 200

601 GAGCCCCAGAGTGTGCCACTTCTGGCCAGATCCCCAAGCACCAACAGGAAATACCCACCC 660
201 E P Q S V P L L A R S P S T N R K Y P P 220

661 CTGCCCCGTGACAAGCTGGAAGAGGAAATTAACCGGAGAATGGCAGACGACAATAAGCTC 720
221 L P V D K L E E E I N R R M A D D N K L 240

721 TTCAGGGAGGAATTCAACGCTCTCCCTGCATGTCCTATCCAGGCCACCTGTGAGGCTGCT 780
241 F R E E F N A L P A C P I Q A T C E A A 260

781 TCCAAGGAGGAAAACAAGGAAAAAATCGATATGTAAACATCTTGCCTTATGACCACTCT 840
261 S K E E [N K E K N R Y V N I L P Y D H S 280
PTPase DOMAIN I

FIG.8A

841 AGAGTCCACCTGACACCGGTTGAAGGGTTCCAGATTCTGATTACATCAATGCTTCATTC 900
281 R V H L T P V E G V P D S D Y I N A S F 300

901 ATCAACGGTTACCAAGAAAAGAACAAATTCATTGCTGCACAAGGACCAAAAAGAAGAAACC 960
301 I N G Y Q E K N K F I A A Q G P K E E T 320

961 GTGAATGATTTCTGGCGGATGATCTGGGAACAAAACACAGCCACCATCGTCATCGTTACC 1020
321 V N D F W R M I W E Q N T A T I V M V T 340

1021 AACCTGAAGGAGAGAAAGGAGTGCAAGTGGCCCCAGTACTGGCCAGACCAAGGCTGCTGG 1080
341 N L K E R K E C K C A Q Y W P D Q G C W 360

1081 ACCTATGGGAATATTCGGGTGTCTGTAGAGGATGTGACTGTCCTGGTGGACTACACAGTA 1140
361 T Y G N I R V S V E D V T V L V D Y T V 380

1141 CGGAAGTTCTGCATCCAGCAGGTGGGCGACATGACCAACAGAAAGCCACAGCGCCTCATC 1200
381 R K F C I Q Q V G D M T N R K P Q R L I 400

1201 ACTCAGTTCCACTTTACCAGCTGGCCAGACTTTGGGGTGCCTTTTACCCCGATCGGCATG 1260
401 T Q F H F T S W P D F G V P F T P I G M 420

1261 CTCAAGTTCCTCAAGAAGGTGAAGGCCTGTAACCCTCAGTATGCAGGGGCCATCGTGGTC 1320
421 L K F L K K V K A C N P Q Y A G A I V V 440

1321 CACTGCAGTGCAGGTGTAGGGCGTACAGGTACCTTTGTCGTCATTGATGCCATGCTGGAC 1380
441 H C S A G V G R T G T F V V I D A M L D 460

1381 ATGATGCATACAGAACGGAAGGTGGACGTGTATGCCTTTGTGAGCCGGATCCGGGCACAG 1440
461 M M H T E R K V D V Y G F V S R I R A Q 480

1441 CGCTGCCAGATGGTGCAAACCGATATGCAGTATGTCTTCATATACCAAGCCCTTCTGGAG 1500
481 R C Q M V Q T D M Q Y V F I Y Q A L L E] 500

1501 CATTATCTCTATGGAGATACAGAACTGGAAGTGACCTCTCTAGAAACCCACCTGCAGAAA 1560
501 H Y L Y G D T E L E V T S L E T H L Q K 520

1561 ATTTACAACAAAATCCCAGGGACCAGCAACAATGGATTAGAGGAGGAGTTTAAGAAGTTA 1620
521 I Y N K I P G T S N N G L E E E F K K L 540

FIG.8B

1621 ACATCAATCAAAATCCAGAATGACAAGATGCGGACTGGAAACCTTCCAGCCAACATGAAG 1680
 541 T S I K I Q N D K M R T G N L P A [N M K 560
 PTPase Domain II

1681 AAGAACCGTGTTTTACAGATCATTCCATATGAATTCAACAGAGTGATCATTCCAGTTAAG 1740
 561 K N R V L Q I I P Y E F N R V I I P V K 580

1741 CGGGGCGAAGAGAATACAGACTATGTGAACGCATCCTTTATTGATGGCTACCGGCAGAAG 1800
 581 R G E E N T D Y V N A S F I D G Y R Q K 600

1801 GACTCCTATATCGCCAGCCAGGGCCCTCTTCTCCACACAATTGAGGACTTCTGGCGAATG 1860
 601 D S Y I A S Q G P L L H T I E D F W R M 620

1861 ATCTGGGAGTGGAATCCTGCTCTATCGTGATGCTAACAGAACTGGAGGAGAGAGGCCAG 1920
 621 I W E W K S C S I V M L T E L E E R G Q 640

1921 GAGAAGTGTGCCAGTACTGGCCATCTGATGGACTGGTGTCTATGGAGATATTACAGTG 1980
 641 E K C A Q Y W P S D G L V S Y G D I T V 660

1981 GAACTGAAGAAGGAGGAGGAATGTGAGAGCTACACCGTCCGAGACCTCCTGGTCACCAAC 2040
 661 E L K K E E E C E S Y T V R D L L V T N 680

2041 ACCAGGGAGAATAAGAGCCGGCAGATCCGGCAGTTCCACTTCCATGGCTGGCCTGAAGTG 2100
 681 T R E N K S R Q I R Q F H F H G W P E V 700

2101 GGCATCCCCAGTGACGGAAGGGCATGATCAGCATCATCGCCGCCGTGCAGAAGCAGCAG 2160
 701 G I P S D G K G M I S I I A A V Q K Q Q 720

2161 CAGCAGTCAGGGAACGACCCCATCACCGTGCAGTGCAGCGCCGGGCGAGGAAGGACGGGG 2220
 721 Q Q S G N H P I T V H C S A G A G R T G 740

2221 ACCTTCTGTGCCCTGAGCACCGTCCTGGAGCGTGTGAAAGCAGAGGGGATTTTGGATGTC 2280
 741 T F C A L S T V L E R V K A E G I L D V 760

2281 TTCCAGACTGTCAAGAGCCTGCGGCTACAGAGGCCACACATGGTCCAGACACTGGAACAG 2340
 761 F Q T V K S L R L Q R P H M V Q T L E Q 780

2341 TATGAGTTCTGCTACAAGGTGGTGCAGGAGTATATTGATGCATTCTCAGATTATGCCAAC 2400
 781 Y E F C Y K V V Q E] Y I D A F S D Y A N 800

2401 TTCAAGTAA 2409
 801 F K * 803

FIG.8C